

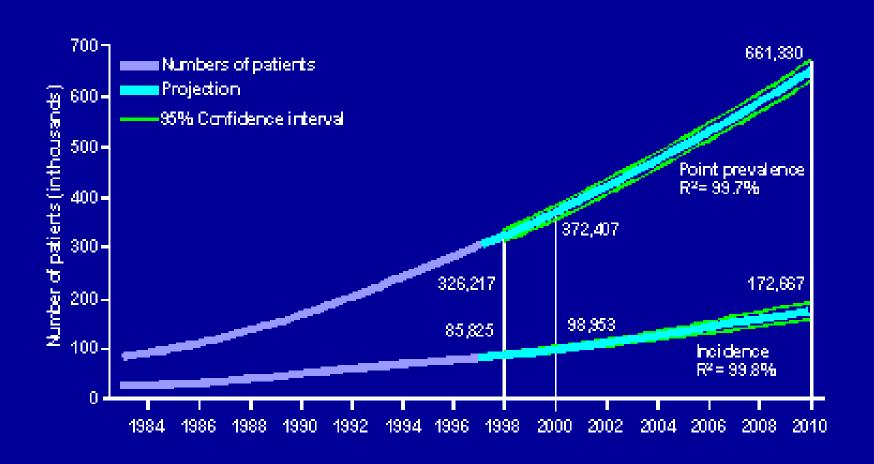
Diabetes, Hypertension, and More:

Managing the Patient with Chronic Kidney Disease (CKD)

Presented by the



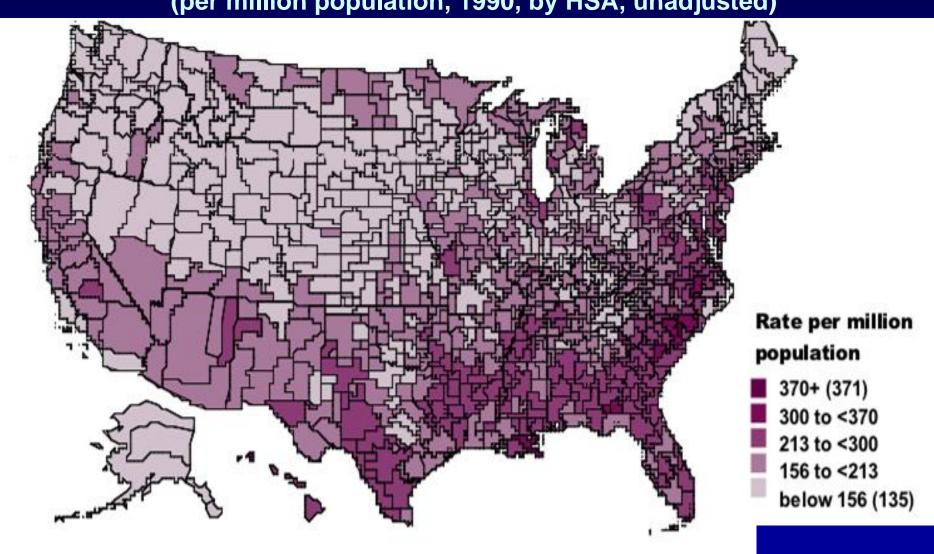
Trends in Incidence and Prevalence of ESRD



US Renal Data System.
USRDS 2000 Annual Data Report.

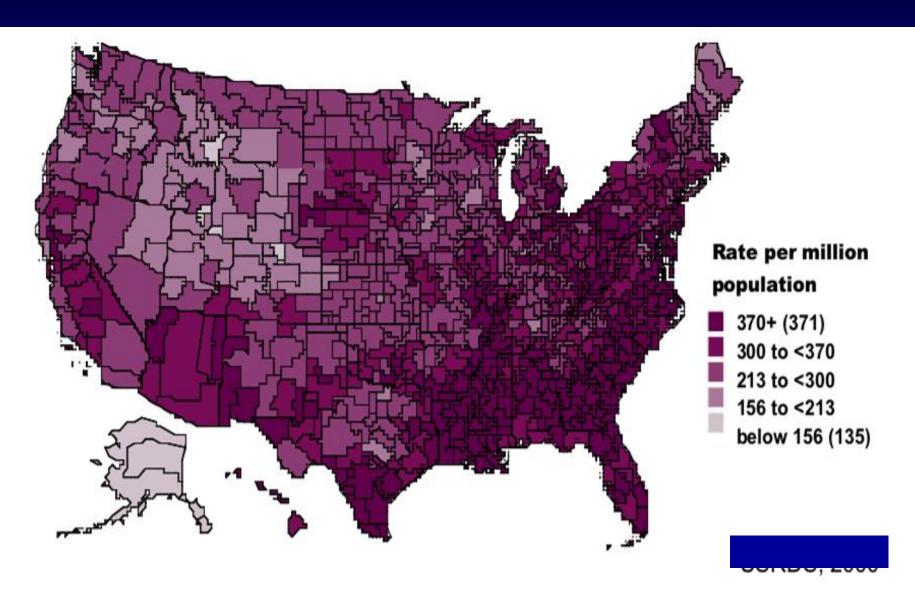
Incidence of Kidney Failure

(per million population, 1990, by HSA, unadjusted)



Incidence of Kidney Failure

(per million population, 2000, by HSA, unadjusted)



The Human Burden

 More than <u>500,000</u> Americans require dialysis or a kidney transplant to stay alive (<u>1,542 patients per million population</u>)

SOUTH CAROLINA:

- About 6700 patients are on dialysis in SC.
- The incidence rate is over 400 patients per million population
- In the last six years, the number of patients currently on dialysis and new cases has **increased by over 30%**
- SC is, per capita, third in the nation with the number of patients on dialysis

The Financial Burden

- Nearly 23 <u>billion</u> Medicare dollars were spent to treat patients with kidney failure in 2006 (and 13.3 billion non-Medicare \$)
- Medicare spending for kidney failure has been increasing at 5-10% per year, based primarily on growing number of patients
- Kidney failure patients constitute 1.2% of the Medicare population but require approximately 6.4 of the expenditures

The Financial Burden

- Hemodialysis costs \$ 71,889/ year (Medicare)
- Peritoneal dialysis \$53,327 / year
- Renal transplant \$ 24,952 / year
- Medicare costs, per patient per month

ESRD \$4900 CKD \$600

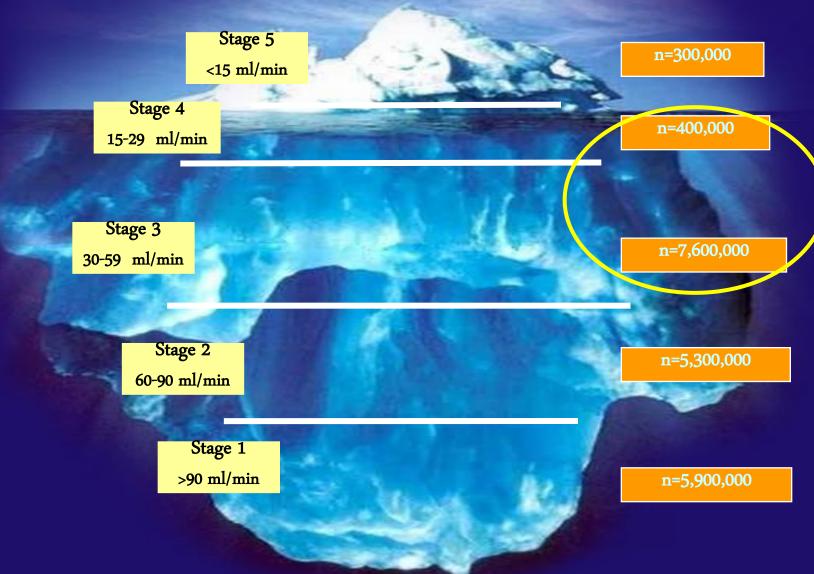
Life Expectancy for Selected Diseases



USRDS 1993.

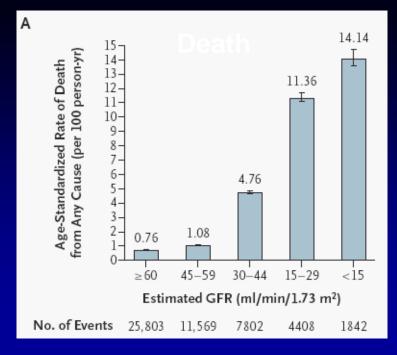


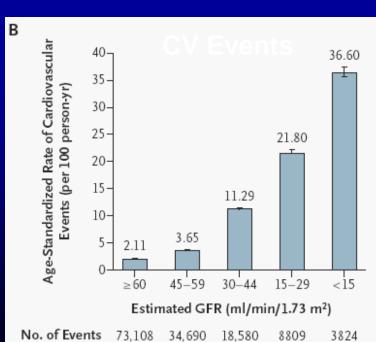
CKD population of 20M in U.S.



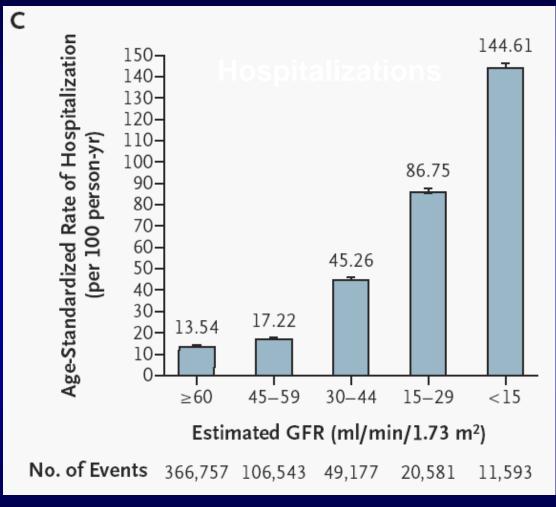
The Problem of Chronic Kidney Disease (CKD)

- 1 in 8 South Carolinians or 460, 000 residents have chronic kidney disease
- 75% of kidney patients in SC are African American
- Hospitalization rates are three times higher in CKD patients
- More than \$121 million was charged in 2004 for hospital visits in SC for kidney disease patients (SC DHEC)
- Nationally, patients with CKD account for 19.4% of Medicare costs in the year the disease is identified. (6.6% of the population)





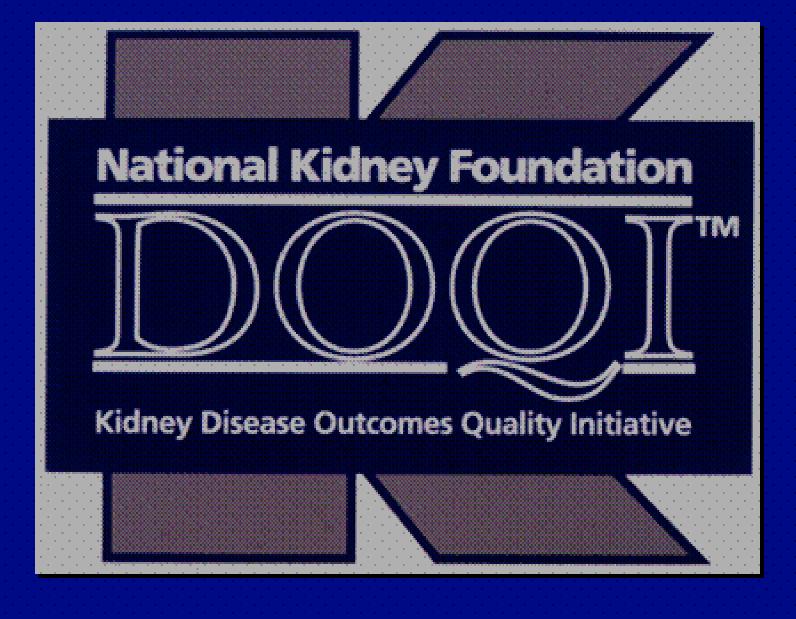
Economic Impact of ↓ GFR Seen in Early Stages of CKD



Go, et al. *N Engl J Med*. 2004;351:1296-1305

Chronic Kidney Disease – The Facts

- Common & incidence is on the rise!!!
- Serious
- Costly
- Under-diagnosed & under- treated
- Preventable
- Manageable



Definition of CKD

- Kidney damage for ≥3 months
 - Defined by structural or functional abnormalities of the kidney, with or without decreased glomerular filtration rate (GFR)

• GFR <60 mL/min/1.73 m² for ≥3 months

 New staging for CKD is primarily based on kidney function

National Kidney Foundation – Kidney Disease Outcomes Quality Initiative (NKF-K/DOQI) Stages of Chronic Kidney Disease

Stage	Description	GFR (ml/min/1.73 m²)
1	Kidney Damage with Normal or ↑ GFR	>90
2	Mild ↓ GFR	60-89
3	Moderate ↓ GFR	30-59
4	Severe ↓ GFR	15-29
5	Kidney Failure	<15 or Dialysis

Why Estimate GFR From SCr, Instead of Using SCr for Kidney Function?

Age	Gender	Race	SCr (mg/dL)	eGFR (mL/min/1.73 m²)	CKD Stage
20	M	В	1.3	91	1*
20	M	W	1.3	75	2 *
55	M	W	1.3	61	2 *
20	F	W	1.3	56	3
55	F	В	1.3	55	3
50	F	W	1.3	46	3

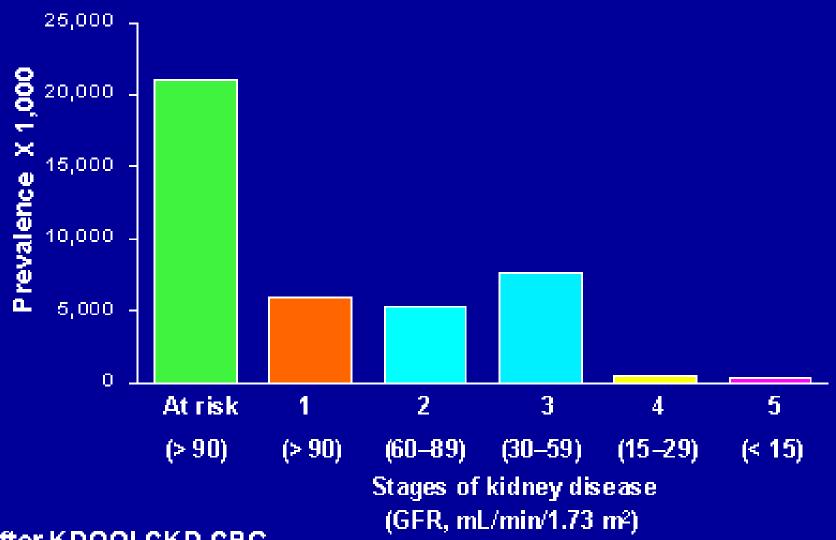
B = black; W = all ethnic groups other than black; *With evidence of kidney damage.

The Prevalence of CKD is High

NKF-KIDOQI Estimates of Prevalence of CKD in the U.S.

		GFR	Prevalen	Prevalenc
Stage	Description	(ml/min/1.73	ce	е
		m²)	(000s)	(%)
1	Kidney Damage with Normal or ↑ GFR	>90	5900	5.8
2	Mild ↓ GFR	60-89	5300	3.0
3	Moderate ↓ GFR	30-59	7600	4.3
4	Severe ↓ GFR	15-29	400	0.2
5	Kidney Failure	<15 or	300	0.1
		Dialysis		

Stages And Prevalence Of Chronic Kidney Disease



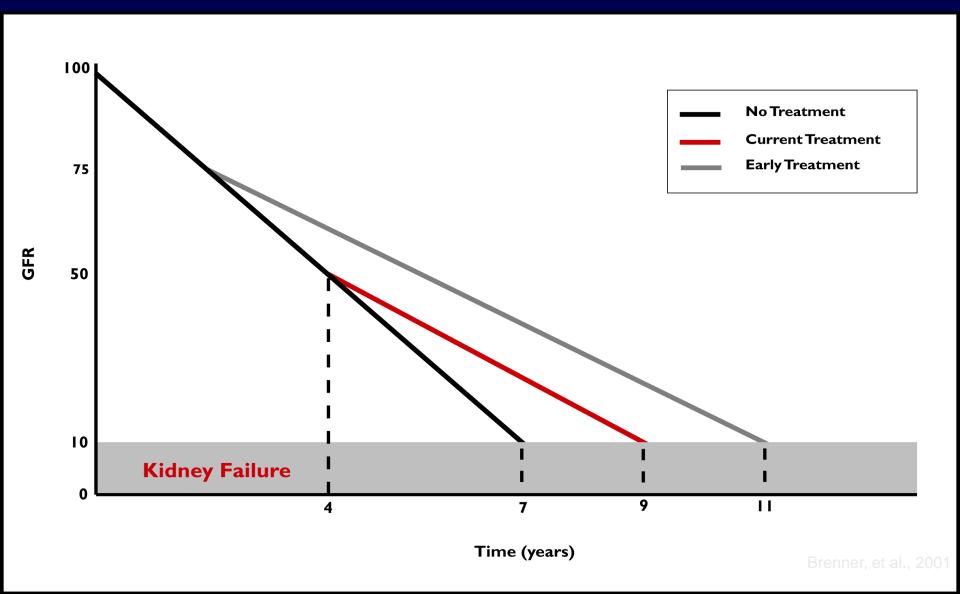
After KDOQI CKD CPG.

Summary Clinical Action Plan for Chronic Kidney Disease

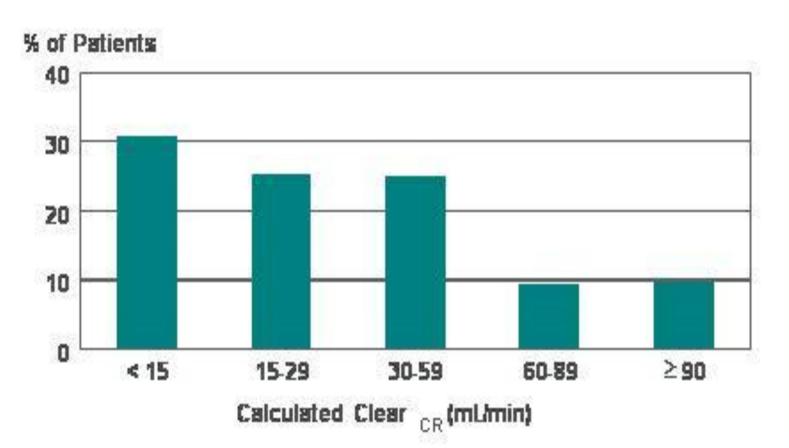
Stage	Description	GFR (ml/min/1.73 m²)	Action Plan (build at each stage)
	At Increased Risk	>90 (CKD Risk Factors)	Screening, CKD Risk Reduction
1	Kidney Damage with Normal or ↑ GFR	>90	Diagnosis and Treatment, Slowing Progression CVD Risk Reduction
2	Mild ↓ GFR	60-89	Estimating Progression
3	Moderate ↓ GFR	30-59	Evaluating and Treating Complications
4	Severe ↓ GFR	15-29	Preparation for Kidney Replacement Therapy
5	Kidney Failure	<15 or Dialysis	Replacement, if Uraemia Present

Adapted from Am J Kid Dis 2002: 39 (No 2, suppl.1): \$17-\$31

Early Treatment Makes a Difference

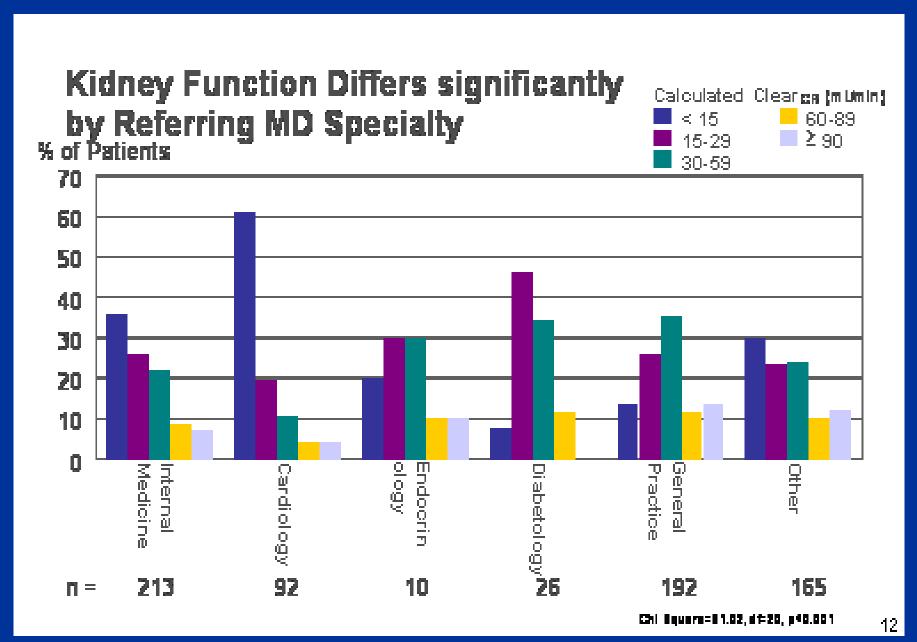


Kidney Function at Time of Referral



CKD Is Not Adequately Recognized and Not Treated Early

- Most practices screen fewer than 10% of their Medicare patients who have diabetes, the single biggest risk factor for CKD
- Patients are referred late to a nephrologist, especially African-American men, who are at high risk for CKD
- Less than 1/3 of people with identified CKD get an ACE Inhibitor to control their blood pressure—and high blood pressure is a major risk factor for CKD
- Nearly 50% of HD patients do not receive Vitamin D replacement therapy, and the Vitamin D use in Stage 3-4 is well under 50%, despite the known protective effects of Vitamin D in kidney disease



EVALUATION OF CKD

Evaluation Of CKD: Causes

- Diabetes mellitus ~50%
 look for microalbuminuria, retinopathy, long history
- Hypertension ~30%
 look for poor BP control, family history of CKD, proteinuria but never nephrotic syndrome
- Glomerular Disease hematuria, RBC casts, nephrotic range proteinuria
- Obstruction
 BPH symptoms; Gyn or GI cancers, hematuria, freq. UTI's
- Medications interstitial nephritis and other disorders

Which Patients Should Be Screened for CKD?

Susceptibility Risk Factors	Progression Factors
 Diabetes Hypertension Older age Family history of nephropathy Racial or ethnic minority Other Low income/minimal education Reduction in kidney mass Known kidney disease 	 Higher level of proteinuria Higher BP Poor glycemic control Smoking Hyperlipidemia Drug use

Evaluation Of CKD: Laboratory workup

For all patients with CKD or increased risk for CKD, evaluate

- Estimated GFR (eGFR)*
 - Calculate annually from serum creatinine (SCr), age, gender, and race
 - Automatic calculator available at: www.kidney.org
- Urine examination
 - Examine the urine sediment or dipstick for erythrocytes and leukocytes
 - Albumin:creatinine ratio in random, untimed urine specimen
 - Microalbuminuria is 30-299 mg Alb/g Cr
 - Clinical albuminuria is ≥300 mg Alb/g Cr
 - Begin screening at the time of type 2 diabetes diagnosis
 - Begin screening at 5 years for type 1 diabetes

GFR: Measurement vs Estimate

Measured 24-hour urine collection for creatinine Clearance

- Formulas for estimating GFR
 - Cockcroft-Gault

mL/min =
$$72 \times SCr \times 0.85$$
 if female

MDRD (A PDA version can be obtained at this NKF web site)

www.kidney.org/professionals/kdoqi/cap.cfm

```
mL/min/1.73 m<sup>2</sup> = 170 × (SCr)<sup>-0.999</sup> × (age)<sup>-0.176</sup> × (BUN)<sup>-0.170</sup> × (Alb)<sup>0.318</sup> × (0.762 if female) × (1.180 if black)
```

Evaluation of CKD

- Estimate GFR with MDRD calculation, urinalysis and micro, spot urine for protein/creat ratio or microalbumin/creat ratio if diabetic
- Renal ultrasound
 - Two kidneys; no obstruction; no masses
 - Asymmetric size suggests renovascular disease
- Is patient over age 50 or very anemic?
 - Check SPEP, UPEP to r/o multiple myeloma
- Is renal function stable?
 - Frequent creatinine check, until stability clear

Evaluation Of CKD

Refer to a nephrologist if:

- you are uncomfortable with diagnosis
- diagnosis may be a GN
 - ... > 3 grams protein in a non-diabetic
 - ... an active urine sediment

renal function declining rapidly

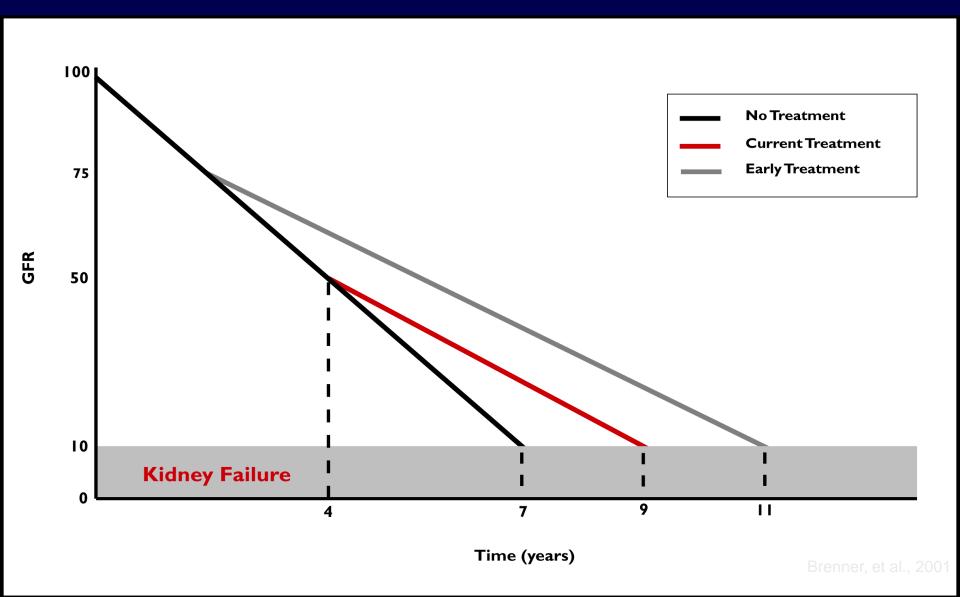
MANAGEMENT OF EARLY CKD

STAGE 1 & 2

Management Of CKD (Stage 1 & 2)

- eGFR: normal to 60 ml/min
- Identify presence and cause of CKD
- Risk factors for progression
 - Hypertension
 - Proteinuria
 - Poor glucose control
- Assess and address cardiovascular risk factors
 - Lipids & Smoking cessation

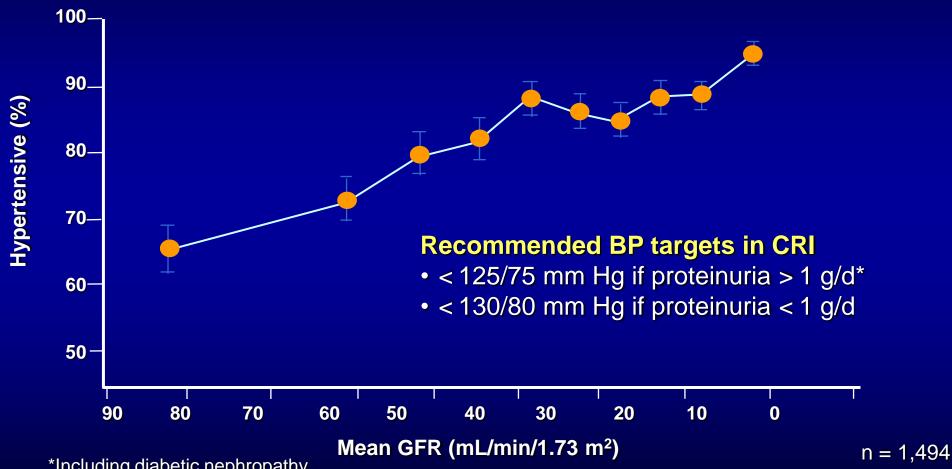
Early Treatment Makes a Difference



Management Of CKD

Hypertension

Hypertension Develops Early And Progresses In CKD

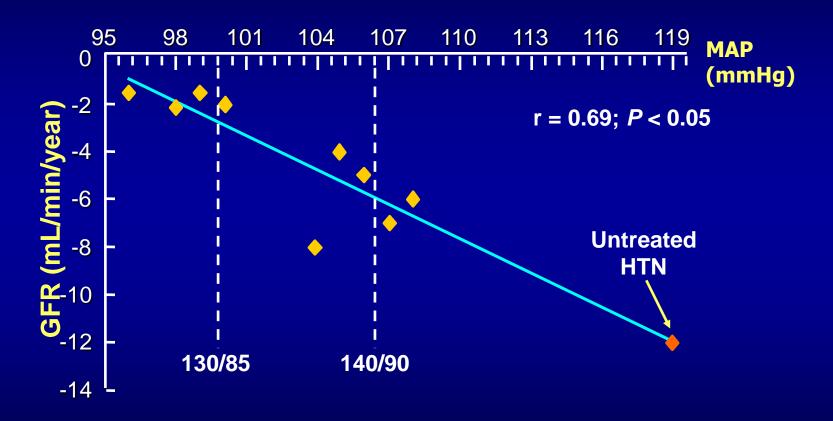


*Including diabetic nephropathy.

Buckalew et al. *Am J Kidney Dis.* 1996;28:811-821.

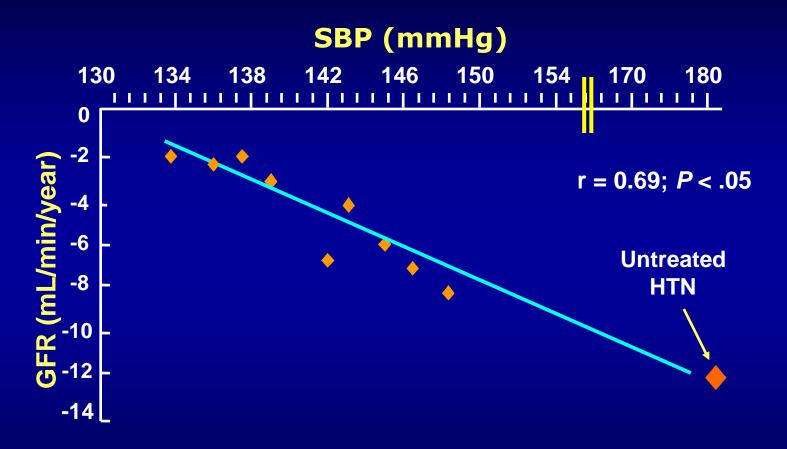
Levey et al. Am J Kidney Dis. 1998; 32:853-906.

Meta Analysis: Mean BP Achieved And Rate Of Decline In GFR In Diabetics And Nondiabetics



Bakris GL, et al. Am J Kidney Dis. 2000;36(3):646-661. Reprinted by permission, Harcourt Inc.

Meta Analysis: Systolic BP Achieved And Rate Of Decline In GFR In Diabetics And Non-diabetics



Bakris GL, et al. Am J Kidney Dis. 2000;36(3):646-661.

Drugs for hypertension

- ACE inhibitors, especially if proteinuria
- Diuretics
- ACE inihibitor/diuretic combination
- Angiotension receptor blocker, especially if proteinuria
- Calcium channel blockers
 - Non-DHP preferable if proteinuria present
 - DHP
- Beta blockers
- Others

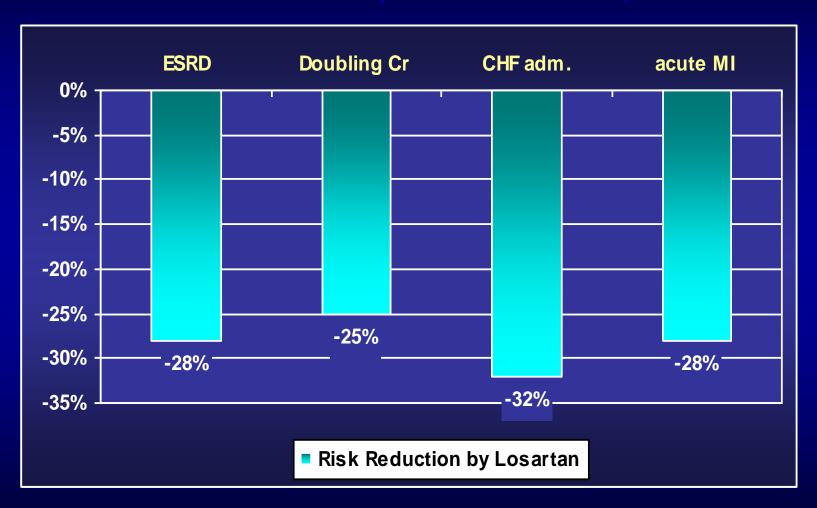
ARB's In Type II Diabetic Nephropathy RENAAL (Losartan Trial)

- Multinational, randomized, double-blinded, placebo-controlled study in 1513 type 2 diabetics with nephropathy (Cr 1.5 – 3.0)
- Primary hypothesis: an ARB with conventional BP meds delays ESRD, or death more than placebo with conventional BP meds



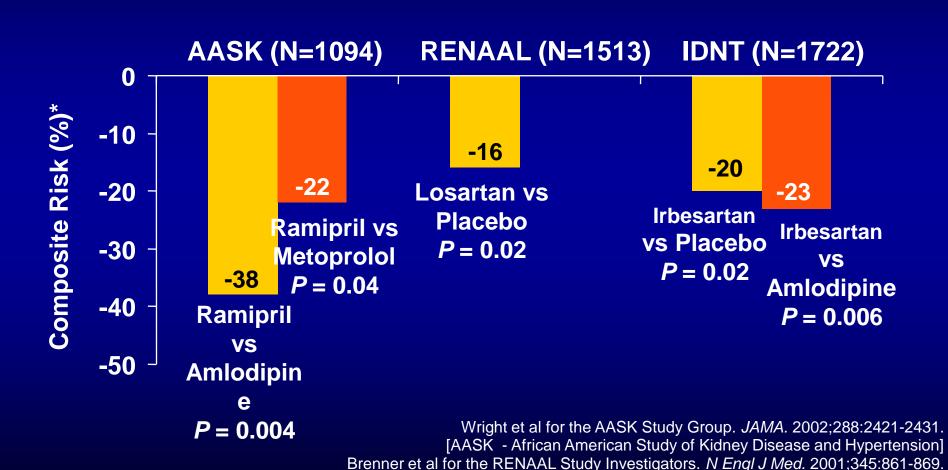
Average follow-up of 3.4 years

ARB's in Type II Diabetic Nephropathy RENAAL (Losartan trial)



BP control was not significantly different between the groups

ACEI/ARB & Reduced Risk of Rapid GFR Decline, Kidney Failure, or Death



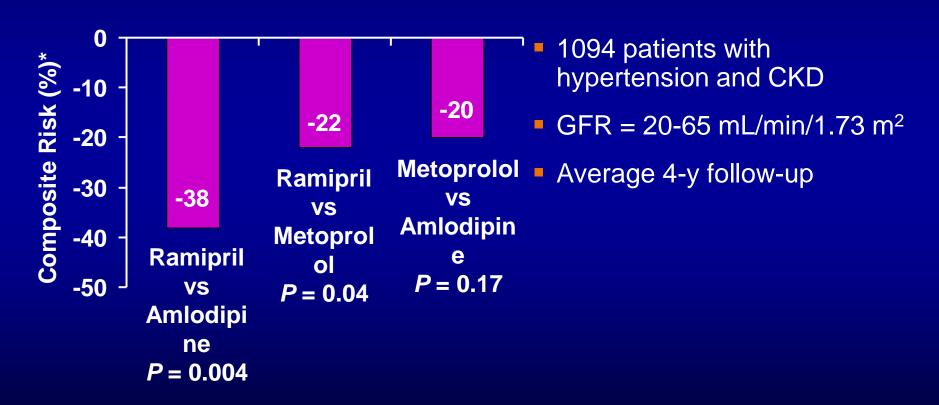
[RENAAL = Reduction of Endpoints in NIDDM with the Angiotensin II Antagonist Losartan]

Lewis et al for the Collaborative Study Group. N Engl J Med. 2001;345:851-860.

[IDNT = Irbesartan in Diabetic Nephropathy Trial.]

Evidence for Reduced Risk of CKD Progression With ACEI Therapy

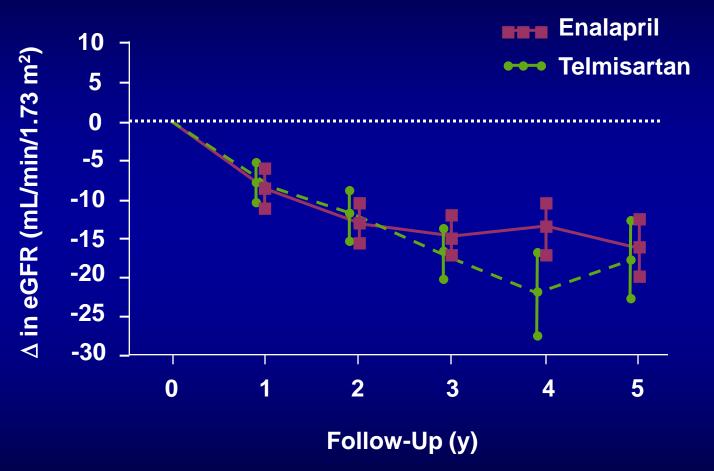
African American Study of Kidney Disease and Hypertension (AASK)



ACEI = angiotensin-converting enzyme inhibitor.*Composite risk of rapid GFR decline/decrease from baseline of 50% or 25 mL/min/1.73 m², kidney failure, or death in patients with existing kidney damage.

Wright et al for the AASK Study Group. *JAMA*. 2002;288:2421-2431.

ACEIs or ARBs Effective for Patients With Type 2 DM

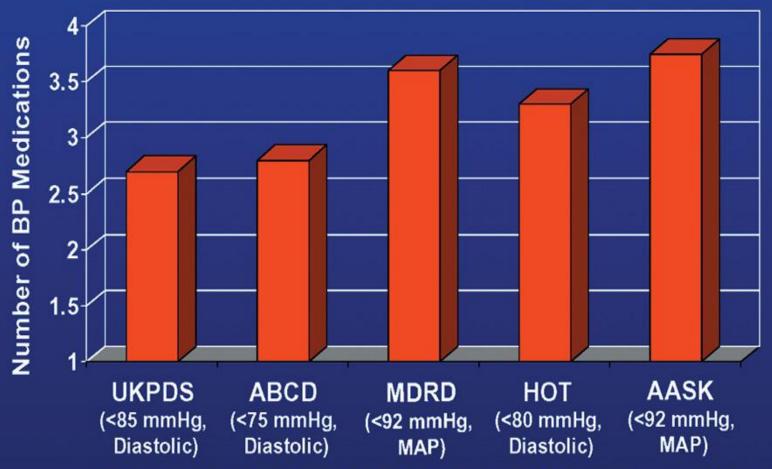


N = 250. Changes from baseline eGFR, based on 5-y data according to treatment group of enalapril or telmisartan in patients with type 2 diabetes and nephropathy.

ARB = angiotensin II receptor blocker.

Barnett et al. N Engl J Med. 2004;351:1952-1961.

Number of Antihypertensive Agents Used to Achieve Target BP



UKPDS=United Kingdom Prospective Diabetes Study, ABCD=Appropriate Blood Pressure Control in Diabetes, MDRD=Modification of Diet in Renal Disease, HOT=Hypertension Optimal Treatment, AASK=African-American Study of Kidney disease and hypertension.

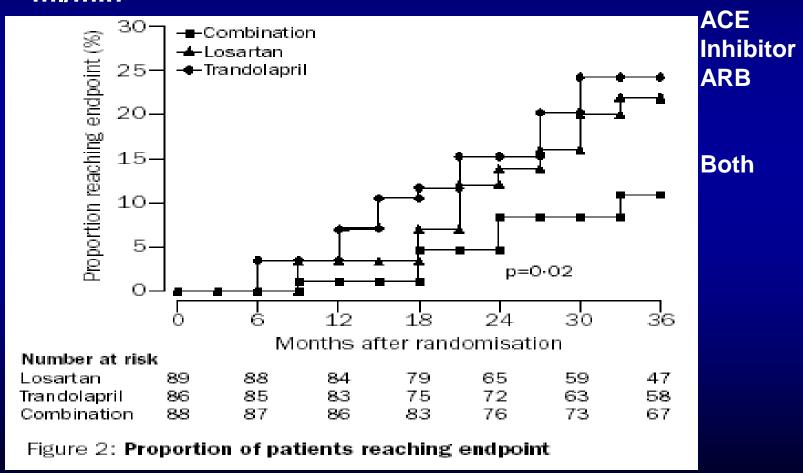
Bakris GL. J Clin Hypertens. 1999;1:141-147.

Management Of CKD

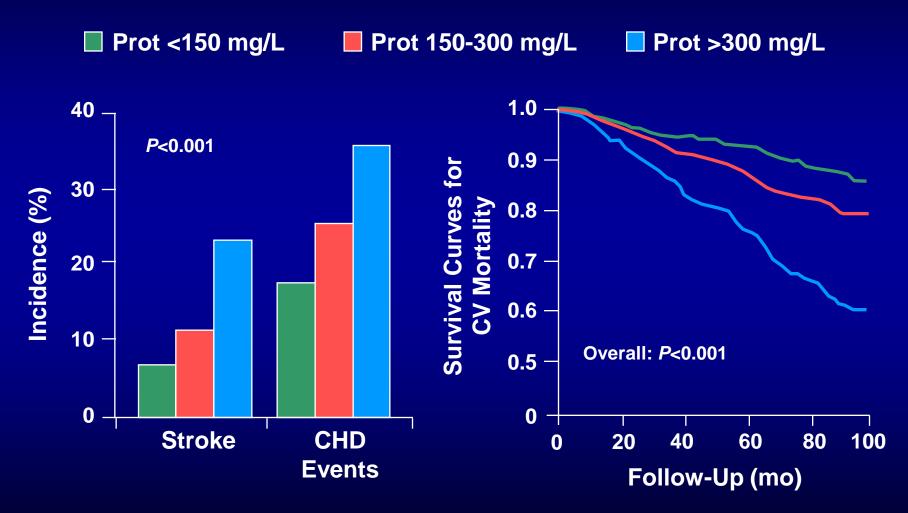
Proteinuria

Management Of CKD: Proteinuria

Combined endpoint: doubling creatinine or est. GFR <7
 ml/min



Proteinuria Predicts Stroke and CHD Events in Patients With Type 2 Diabetes



CHD = coronary heart disease; Prot = urinary protein excretion. Miettinen et al. *Stroke*. 1996;27:2033-2039.

Goals for Glycemic Control in Diabetes

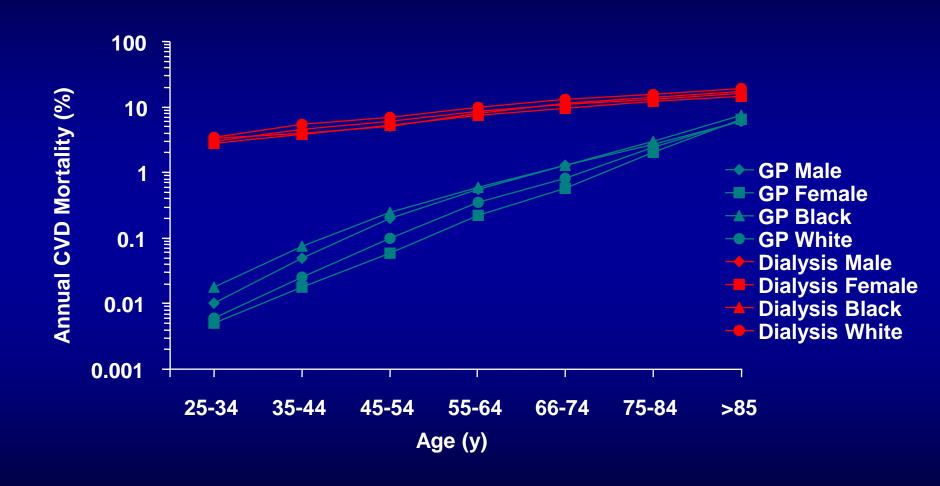
Parameter	Goal		
	ACE/AACE ¹	ADA ²	
A1C	≤6.5% (normal, 4%-6%)	<7%	
Preprandial plasma glucose	<110 mg/dL	90-130 mg/dL	
Postprandial plasma glucose	<140 mg/dL*	<180 mg/dL [†]	

A1C = glycosylated hemoglobin; ACE = American College of Endocrinology; AACE = American Association of Clinical Endocrinologists; ADA = American Diabetes Association.

Management Of CKD

Hyperlipidemia

Cardiovascular Disease Mortality Is Increased in Patients on Dialysis



CVD = cardiovascular disease; CKD = chronic kidney disease; GP = general population. Foley et al. *Am J Kidney Dis.* 1998;32(suppl 3):S112-S119.

Incidence of Cardiovascular Disease (CVD) and Rate of Renal Function Loss in Stages 1 to 3 of CKD

- Compared the incidence rates of CVD in subjects with stages 1-3
- Data from PREVEND study, an observational cohort study performed in the Netherlands

• Median Prevalence and Demographics of CKD and CVD in Stages 1-3

		-O V S				
		No CKD	Stage 1	Stage 2	Stage 3	
	N	6906	243	856	491	
	Age, yr (SD)	47 (12)	48 (12)	56 (12)	63 (9)	
	Male, %	49	66	64	38	
L	Incidence Rates of CVD	7.1	16.9*	22.8*	22.5*	
	Age and sex-adjusted Hazard Ratio for CVD	Ref	2.2 (1.5-3.2)	1.6 (1.3-2.0)	1.3 (1.0-1.7)	

Outcomes Based on Presence of Microalbuminuria (MA)

	Stage 3, MA -	Stage 3, MA +	
Hazard Ratio for CVD	1.0 (0.7-1.5)	1.7(1.2-2.4)	
Annual Change in eGFR during follow-up (mL/min/1.73m2/yr)	+0.16±1.22	-0.50±1.21**	

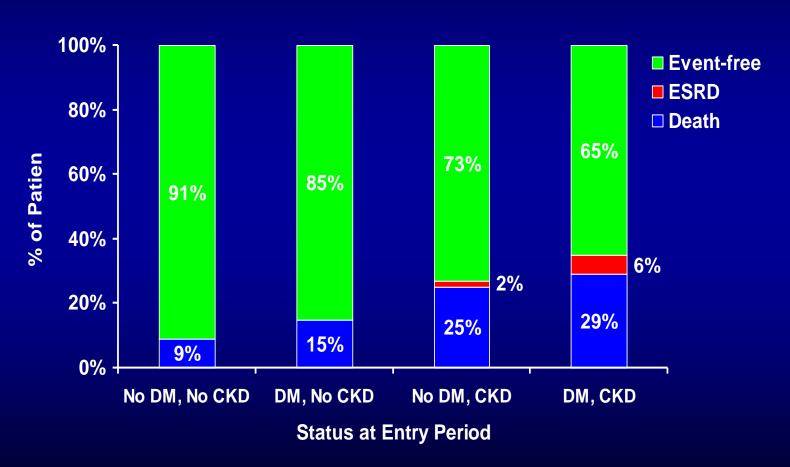
Conclusions:

- Screening for stages 1 and 2 successfully detects subjects at risk for CVD
- For defining stage 3 CKD, an additional criterion determining presence of MA may be needed

*p<0.001, compared to No CKD

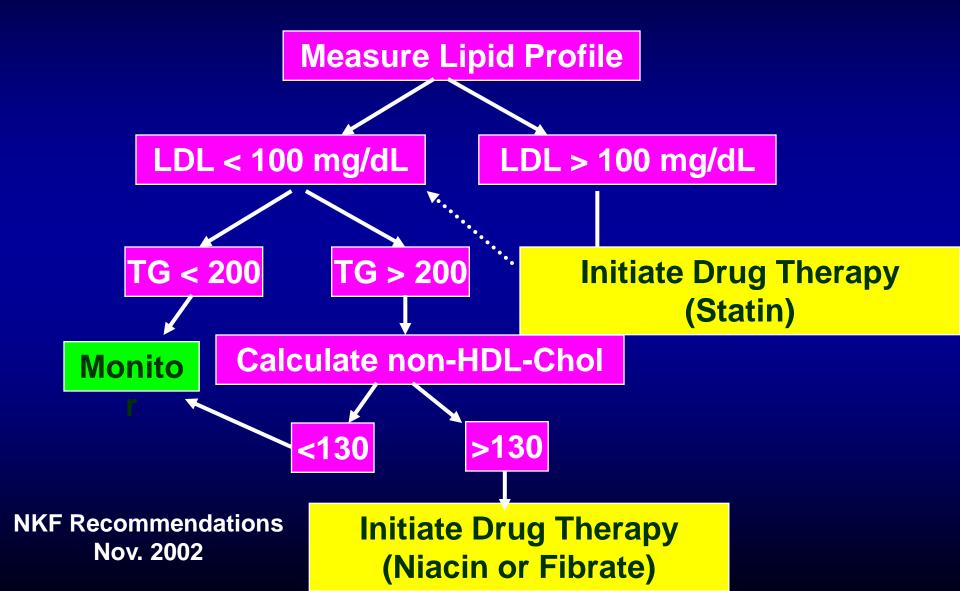
^{**}p<0.001, compared to Stage3, MA-

CKD Patients Are More Likely To <u>Die</u> Than Progress To ESRD Percentage Who Remained Event-free Vs Death Vs Developed ESRD During 2-year Follow-up



Medicare 5% sample 1996-1997, 2-year follow-up, adjusted for age, gender, and race. Analysis performed by Minneapolis Medical Research Foundation.

Lipid Management Targets In CKD



Stage 3 CKD eGFR 30 – 59 ml/min

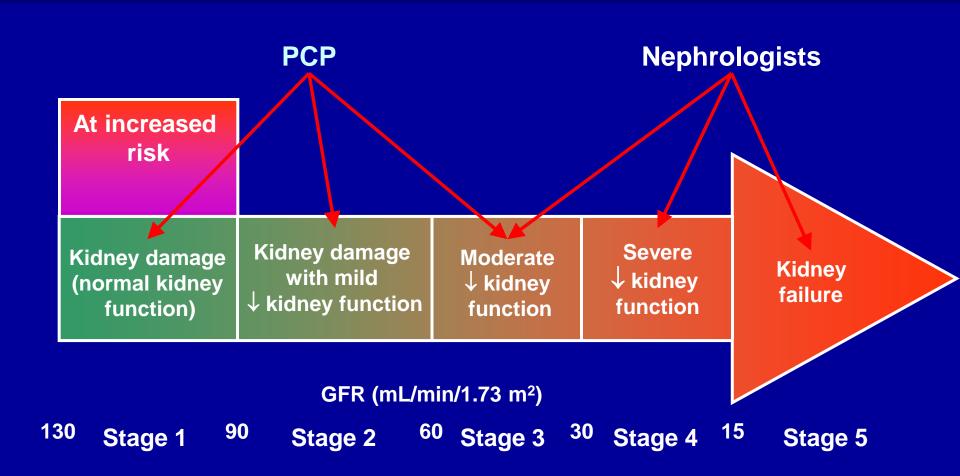
Common Adjustments In Stage 3 CKD GFR 30 To 59 ml/min

- Convert HCTZ to Furosemide to maintain diuresis.
- Stop K+ sparing diuretics.
- Reduce Allopurinol from 300 mg/d to 200 mg/d.
- Stop Glucophage (do not use in stage 2 CKD or Scr >1.5mg/dl)
- Avoid all NSAIDs.
- Assess new drugs for renal dose adjustment.
- Refer all patients to nephrology when GFR below 30 ml/min.

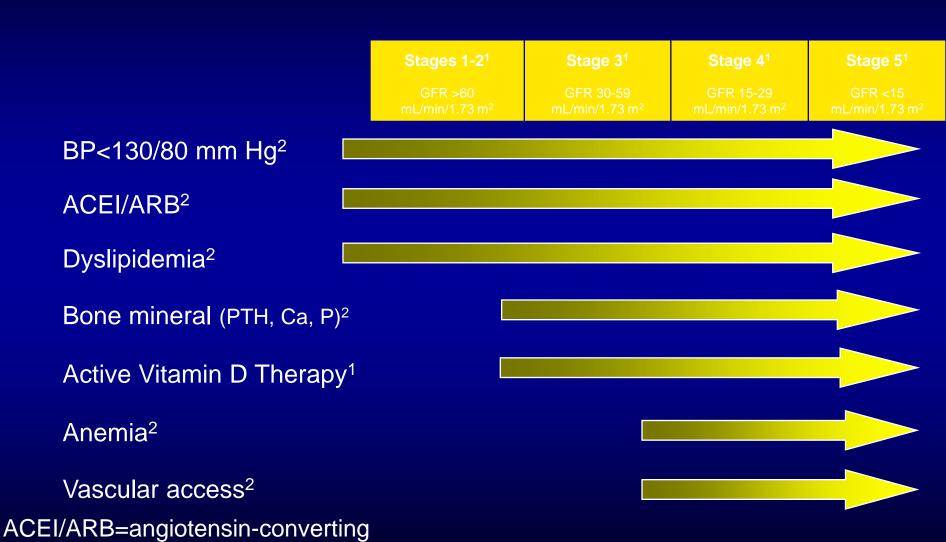
Summary Of Management Objectives

- Screen at-risk population for CKD
- If diabetic, control A_{1c} to < 7%
- Reduce BP to <130/80 mmHg, <125/75 if CKD present
- Use multiple antihypertensive drugs
- Monitor for maximal reduction of proteinuria
- Treat hyperlipidemia (LDL<100mg/dl) & stop smoking
- Enteric coated aspirin 81-325 mg/day
- Low salt diet (NaCl < 2g/day)

Co-Management



CKD Clinical Action Plan



enzyme inhibitor/angiotensin II receptor blocker

1. National Kidney Foundation. *Am J Kidney Dis*. 2003;42:1-201. 2. Modified from Levey et al. *Ann Intern Med*. 2003;139:137-147.

SUMMARY

- CKD is very common
- CKD populations are easily identifiable and accessible
- Guidelines for prevention & management of CKD available
- CKD care can be improved
- CKD patients are at high risk for cardiovascular disease
- Development of ESRD can be delayed

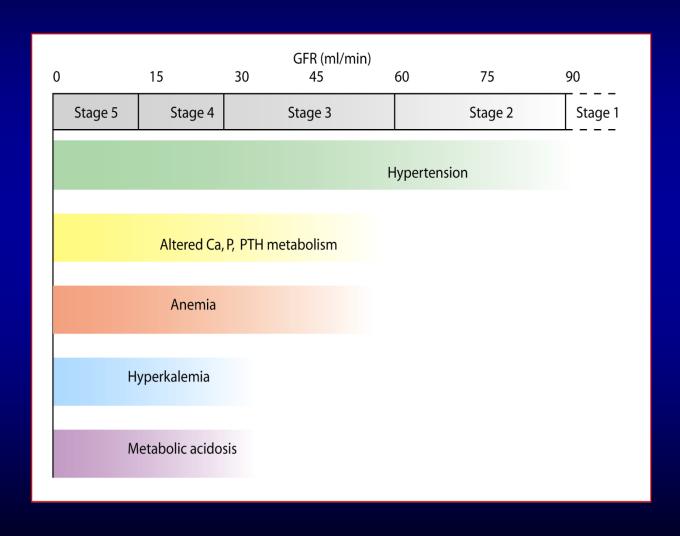
Questions?

Thank you

Dr. Prakash Prabhu ppp0809@hotmail.com

Complications of CKD

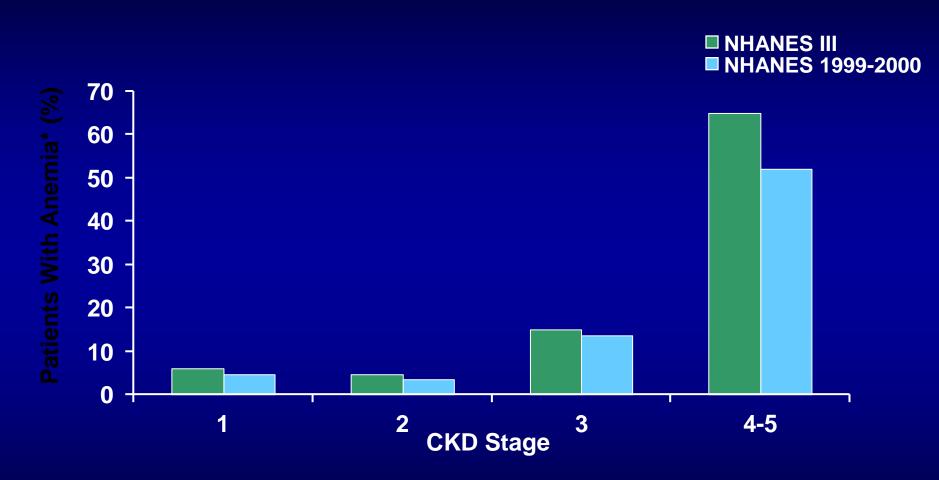
Timing of Complications of CKD



Management of CKD

Anemia

Anemia Prevalence by CKD Stage



*NHANES participants aged ≥20 y with anemia as defined by WHO criteria: hemoglobin (Hgb) <12 g/dL for women, and Hgb <13 g/dL for men.

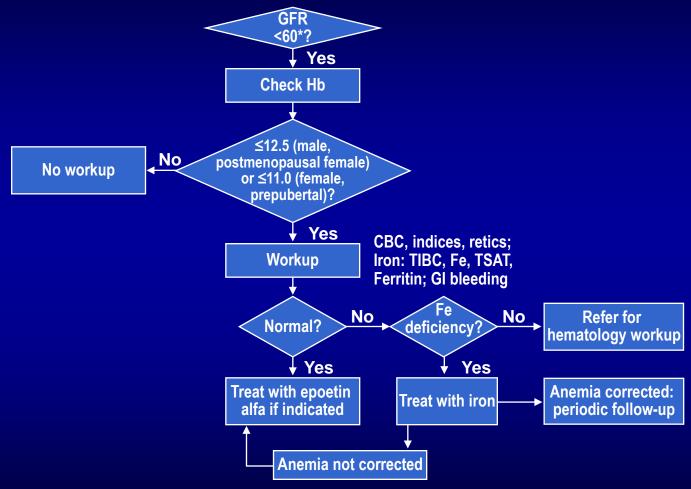
USRDS 2004 Annual Data Report. The data reported here have been supplied by the USRDS. The interpretation and reporting of these data are the responsibility of the author(s) and in no way should be seen as an official policy or interpretation of the U.S. government. Available at: www.usrds.org. Accessed 3/28/05.

Consequences of Anemia in CKD

- Reduced oxygen delivery to tissues
- Decrease in hemoglobin (Hgb) compensated by increased cardiac output
- Progressive cardiac damage and progressive renal damage¹
- Increased mortality risk²
- Reduced quality of life (QOL)³
 - Fatigue
 - Diminished exercise capacity
 - Reduced cognitive function
- Left ventricular hypertrophy (LVH)⁴

^{1.} Silverberg et al. *Blood Purif.* 2003;21:124-130. 2. Collins et al. *Semin Nephrol.* 2000;20:345-349; 3. The US Recombinant Human Erythropoietin Study Group. *Am J Kidney Dis.* 1991;18:50-59; 4. Levin. *Semin Dial.* 2003;16:101-105.

Anemia Assessment Flowchart



CBC = complete blood count; Fe = iron; GFR = glomerular filtration rate; GI = gastrointestinal; TIBC = total iron-binding capacity; TSAT = transferrin saturation.

*mL/min/1.73 m².

Adapted from NKF. Am J Kidney Dis. 2002;39(2 suppl 1):S1-S266.

Early Treatment of Anemia

- Nutritional therapy
 - Fe (daily)
 - Adults: ≥200 mg elemental Fe
 - Children: 2 to 3 mg/kg
 - Folic acid
 - Vitamin B12
- Erythropoietin therapy improves
 - Functional status
 - Quality of life

Target Ranges		
Hgb	11-12 g/dL	
Fe	>20% transferrin saturation >100 ng/mL serum ferritin	

FE = iron.

When and How to Initiate Erythropoietic Therapy in CKD

A Checklist for Starting therapy

- Necessary Laboratory data
 December Creatining and actimated CED at 40 ml/mix
 - ☐ Recent serum Creatinine and estimated GFR < 40 ml/min
 - □ Hemoglobin < 10 g/dl or Hematocrit < 30% (many providers permit treatment when the Hgb < 11 g/dl or Hematocrit < 33%)</p>
- Other required data for billing
 - □ Patient age and weight
 - ☐ Product (Epoetin or Darbepoetin), dose, and frequency
 - ☐ Insurance assessed for coverage method
 - Medicare requires in office administration with MD supervision
 - ☐ Some private insurers require pre-authorization

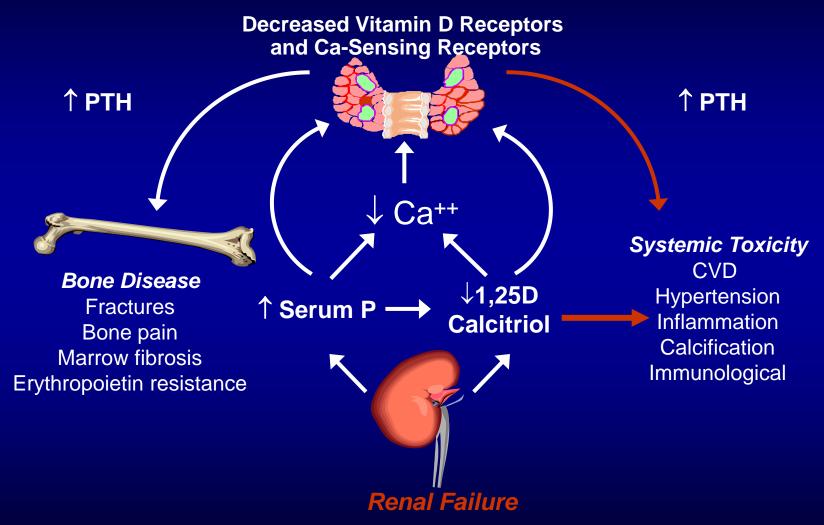
Initial Epoetin and Darbepoetin Dosing

- Epoetin (Procrit): 20,000u multidose vial
 - Starting dose ~ 10,000 u SC weekly or 100 units/Kg weekly
- Darbepoetin (Aranesp): Vials and prefilled syringes
 - Starting dose: 60 mcg vial q 2 weeks; or 40 mcg if < 50 Kg
- Severe anemia can be corrected faster by starting with a higher dose or more frequent dosing, but the increase in Hgb should not exceed 1 g per 2 wks, to avoid hypertension and risk of seizures

Management of CKD

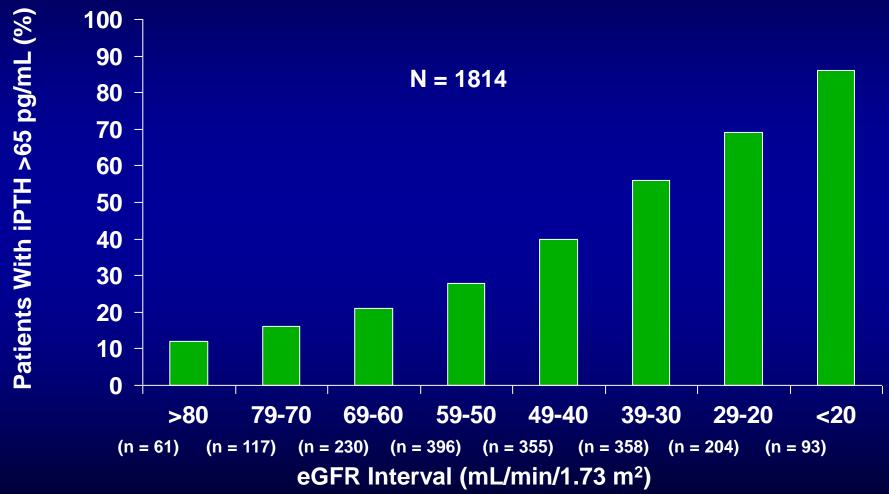
Calcium, Phosphorus & PTH

Feedback Loops in SHPT



Ca = calcium; CVD = cardiovascular disease; P = phosphorus; SHPT = secondary hyperparathyroidism. Courtesy of Kevin Martin, MB, BCh.

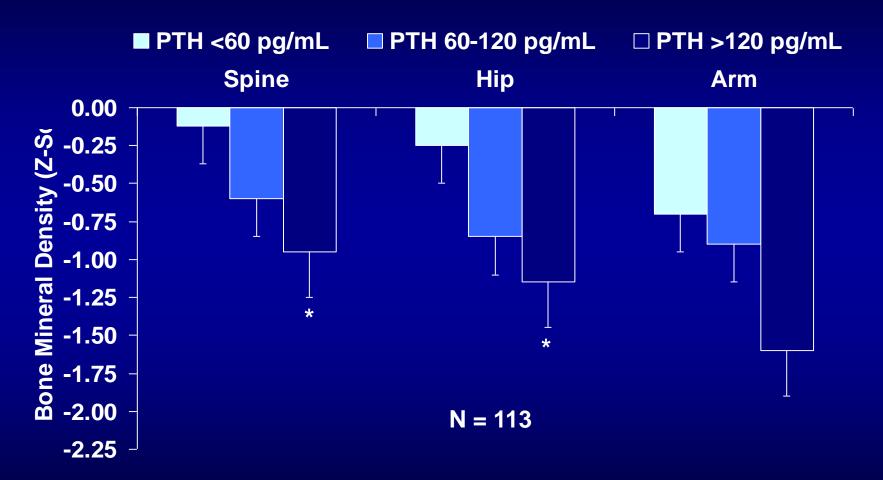
Prevalence of Elevated iPTH by eGFR Intervals



iPTH = intact parathyroid hormone.

Bakris et al. Poster presented at: American Society of Nephrology Renal Week 2005; November 8-13, 2005; Philadelphia, PA. Abstract F-PO732.

Bone Loss Correlates With Severity of SHPT in CKD

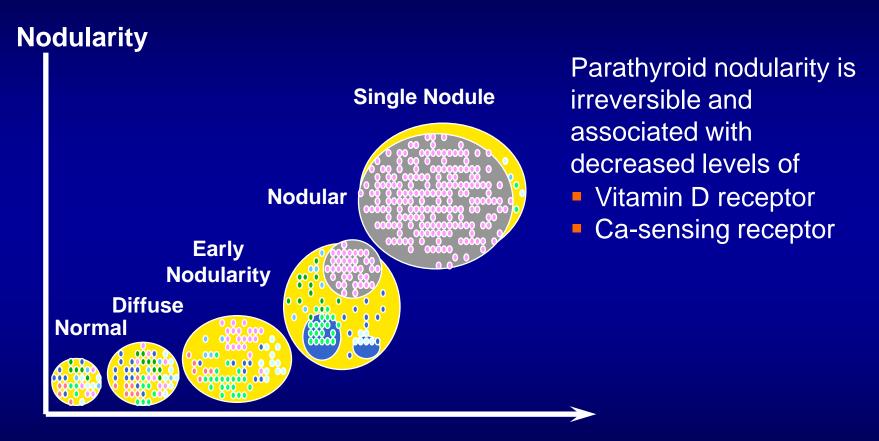


^{*}*P*<0.05 compared with patients with PTH in the normal range.

Z-Score = comparison to the mean value for women at a similar risk, including age, weight and ethnicity.

Rix et al. *Kidney Int.* 1999;56:1084-1093.

The Progression of SHPT in CKD: Parathyroid Growth Becomes Irreversible



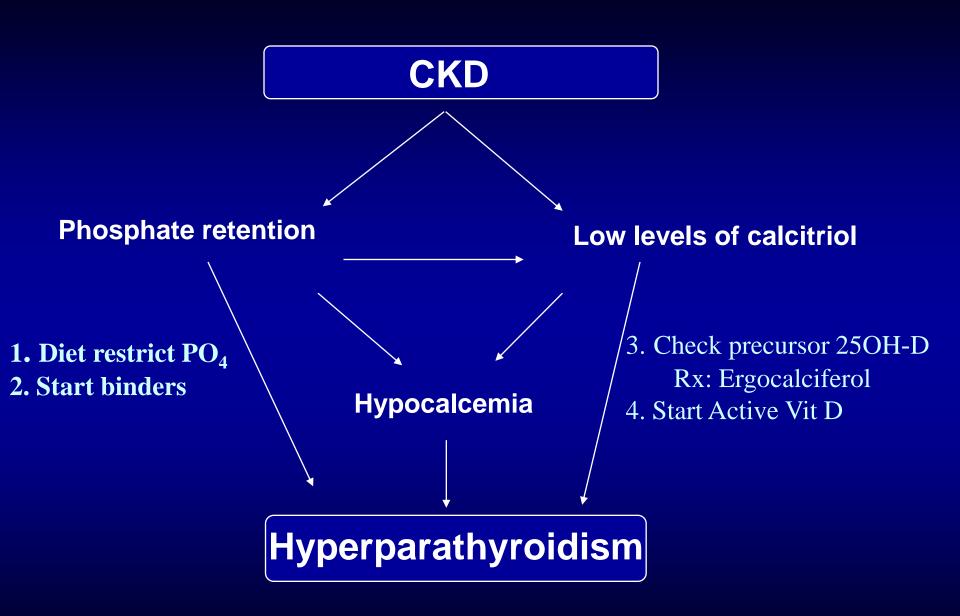
CKD Progression

Early Assessment for Treatable Consequences of Altered Mineral Metabolism in CKD

Consequences in CKD ¹	eGFR (mL/min/1.73 m ²)	CKD Stage
1,25 Vitamin D Deficiency	<65	2-3
Hyperparathyroidism	<65	2-3
Hyperphosphatemia	<40	3-4
Hypocalcemia	<30	4-5

Assessment²

- Stage 3: Ca, P, PTH every 12 mo
- Stage 4: Ca, P, PTH every 3 mo
- If PTH > target range, measure 25(OH)D at first encounter and repeat annually if normal



Monitor Calcium and Phosphorus, PTH at least quarterly

HIGH PHOSPHORUS FOODS

DAIRY

Cheese, all types

Cream

Cream Pies or Desserts

Custard

Frozen Custard

Ice Cream

Ice Milk

Milk, all kinds

Pudding

Yogurt

MISCELLANEOUS

Chocolate

Nuts

BEVERAGES

Beer

Colas

Wyler's Lemonade Mix

PROTEIN FOODS

Braunschweiger

Eggs

Liver

Peanut Butter

Salmon

Sardines

Tuna

VEGETABLES

Bakes Beans and Pork-n-Beans

Beans (Kidney, Lima, Red)

Dried Beans (White, Navy, Pinto)

Dried Peas (Black-eyed, Split Pea)

Lentils

Mixed Vegetables

Soybeans and Soy Foods

BREADS AND CEREAL

Barley

Bran

Cornbread

Waffles (From Mix)

Whole Grain Breads

Phosphate binders

Calcium containing

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Calcium acetate (Phos-Lo)
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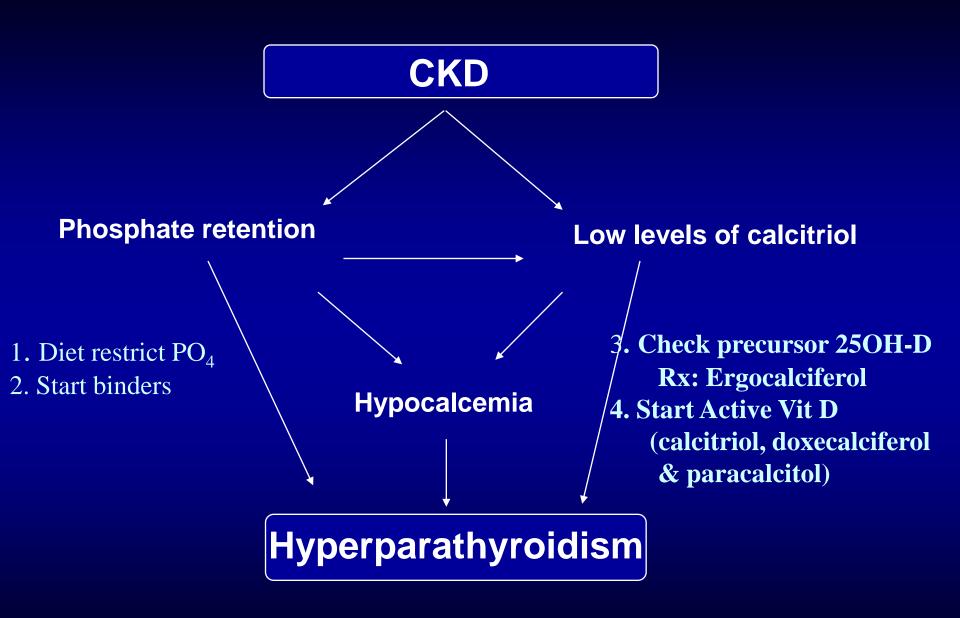
Calcium carbonate (Tums, Tums EX)

Non-calcium containing
 Sevelamer hydrochloride (Renagel)
 Lanthanum carbonate (Fosrenol)

Schedules

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With meals - P binder + Ca supplement
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At bedtime - Ca supplement



Monitor Calcium and Phosphorus, PTH at least quarterly

Vitamin D Metabolism

Active Vitamin D

Inactive D

Liver Kidney
$$1\alpha,25(OH)_2D_3$$

vitamin D_3
 $25(OH)D_3$
 $24,25(OH)_2D_3$

-----Inactive Vitamin D-----

Vitamin D in Chronic Kidney Disease: Stages 3 and 4

Measure serum 25-hydroxyvitamin D in patients with elevated PTH.

If 25-OH-D is normal, repeat annually

Level	Treatment with Ergocalciferol
<5 ng/mL	50,000 U/wk x 12, then q month x 6
5-15 ng/mL	50,000 U/wk x 4, then q month x 6
16-30 ng/mL	50,000 U/month x 6

NKF K/DOQI Guidelines for Bone Metabolism and Disease. Am J Kidney Dis. 2003;42(4 suppl 3):1-201.

Recommended Goals for Hormone and Mineral Metabolism

Parameter	Recommendation
iPTH (pg/mL)	Stage 3: 35-70; stage 4: 70-110
Serum 25(OH) vitamin D (ng/mL)	>30
P (mg/dL)	2.7-4.6
Ca (mg/dL)	Normal parameters for the lab
Ca × P product (mg ² /dL ²)	<55

SUMMARY

- CKD is very common
- CKD populations are easily identifiable and accessible
- Guidelines for prevention & management of CKD available
- CKD care can be improved
- CKD patients are at high risk for cardiovascular disease
- Development of ESRD can be delayed

Questions?

Thank you

Dr. Prakash Prabhu ppp0809@hotmail.com